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<110> Helix Research Institute

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<151> 1999-07-23

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cto	tggg	gact	tgta	gtto	tg g										al Pr	et 293 ro
_										Gln				atg Met 25		341
				-				_	Gly	_		_		gct Ala		389
						_			_	_				ttg Leu	_	437
	_	-									-			caa Gln		485
	-													ctg Leu		533
-														atg Met 105		581
_	-							-						agg Arg		629
age	ctg	gaa	gtg	cct	gag	at.c	aac	gct	t.t.ø	ctg	ctg	et.c	ccc	ลลซ	gag	677

Ser	Leu	Glu 125		Pro	Glu	He	Asn 130	Ala	Leu	Leu	Leu	Val 135	Pro	Lys	Glu	
	tcc Ser 140															725
	gaa Glu															773
	cac His															821
	gag Glu															869
	cag Gln															917
	ggg Gly 220														gtc Val	965
	gga Gly															1013
	acc Thr							Ala								1061
tan.	് മ ന	ga.c	ctø	ot c	cta	202	ort	gra	ወ ወሰ	ลลด	age	tøc	Cgg	gt.ø	gat.	1109

95

Tyr Gln Asp Leu Val Leu Thr Ala Ala Gly Asn Ser Cys Arg Val Asp 270 275 280	
gtc ttc acc aac ctg ggt tac cga gcc ttc tcc ctg tcc ttc Val Phe Thr Asn Leu Gly Tyr Arg Ala Phe Ser Leu Ser Phe 285 290 295	1151
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Ser Lys Lys Pro Val Val Thr Phe Gln Ala His Asp Gly Pro Val Tyr 65 70 75 80	
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90

85

- Glu Lys Ala Trp Leu Trp Ala Glu Met Leu Lys Lys Gly Cys Lys Glu 100 105 110
- Leu Trp Arg Gln Pro Pro Tyr Arg Thr Ser Leu Glu Val Pro Glu
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- Ile Asn Ala Leu Leu Leu Val Pro Lys Glu Asn Ser Leu Ile Leu Ala 130 135 140
- Gly Gly Asp Cys Gln Leu His Thr Met Asp Leu Glu Thr Gly Thr Phe 145 150 155 160
- Thr Arg Val Leu Arg Gly His Thr Asp Tyr Ile His Cys Leu Ala Leu 165 170 175
- Arg Glu Arg Ser Pro Glu Val Leu Ser Gly Gly Glu Asp Gly Ala Val 180 185 190
- Arg Leu Trp Asp Leu Arg Thr Ala Lys Glu Val Gln Thr Ile Glu Val
 195 200 205
- Tyr Lys His Glu Glu Cys Ser Arg Pro His Asn Gly Arg Trp Ile Gly 210 215 220
- Cys Leu Ala Thr Asp Ser Asp Trp Met Val Cys Gly Gly Gly Pro Ala 225 230 235 240
- Leu Thr Leu Trp His Leu Arg Ser Ser Thr Pro Thr Thr Ile Phe Pro 245 250 255
- Ile Arg Ala Pro Gln Lys His Val Thr Phe Tyr Gln Asp Leu Val Leu 260 265 270
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						ctg										706
	Lys	Thr	Pro	Val 5	Glu	Leu	Ala	vai	5er	uly	мес	GIII	1111	15	uly	
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ctt	cag	cac	cgc	tgc	cga	ggt	ggc	tac	cgg	gtc	aag	gcc	agg	acg	tca	754
Leu	Gln	His	Arg	Cys	Arg	Gly	Gly		Arg	Val	Lys	Ala		Thr	Ser	
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						Phe										
		35					40					45				
cca	CCE	gac	ttc	gat	ccg	ccc	tgg	gtg	gag	aag	gct	aac	aga	acc	aga	850
						Pro										
	50					55					60					
						+			++~	~~~	go.	220	aaa	200	t ort	898
						tcg Ser										000
65	Vai	uly	цуз	uiu	70	DCI	Д	Mια	Dou	75		200			80	
						ggc										946
Glu	Thr	Thr	Pro		Arg	Gly	Ser	Thr		Thr	Leu	Thr	Pro		Lys	
				85					90					95		
aag	aac	aaa	tac	aga.	ccc	atc-	age	cac	acc	ccg	tct	tac	tgt	gat	gag.	994
						Ile										
			100					105					110			
t o or	o+ a	+++	aac	tee	റമാ	tct	gaa	ወ ወር	ጀርር	agc	ttc	ggg	gcc	ccg	cgg	1042
															Arg	
D 01	Dou	115	41	501	0		120					125				
																4000
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Pro	Pro	Thr	Pro	Arg		Ser	His	Ser	Pro		Pro	Arg	Glu	Ala		
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Leu	Arg	Ala	116		Pro	Ala	GIY	Pro		Lys	Inr	GIU	Pro	Gly	Pro	
				165					170					175		
gcg	gca	gac	tcc	cag	aag	tta	tct	atg	ggt	ggg	tta	cac	tct	tca	cgc	1234
	_													Ser		
		F	180		_0 -			185	0				190		0	
			100					100								
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Pro	Leu	Lys	Arg	Gly	Leu	Ser	His	Ser	Leu	Thr	His	Leu	Asn	Val	Pro	
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Ser	Thr	Gly	His	Pro	Ala	Thr	Ser	Ala	Pro	His	Thr	Asn	Gly	Pro	Gln	
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														cca		1426
Thr	Ser	Arg			Ser	Val	Ser			Val	Pro	Ser	Thr	Pro	Arg	
				245					250					255		
					~~~				000	a a t	+ ~~	000	t an	tooto	.++	1475
													iga	tacto		1410
Arg	uly	-	Ala	III.	GIII	гуs			Pro	Fro	тгр	LyS				
			260					265								
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Juan	~ <b>u</b> 55	<b>5</b>	B-C-C-C-C	4 455	6 6°	ouce	ovea	. cub	0 040	000	3001		o (	20 246	00000	1000
catt	catc	ac c	cagg	gaac	c cc	aggt.	atta	aag	aagc	ccc	tgtg	ggge	ca i	gacag	acata	1595
				J		-00				-	5-0		•			

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<213> Homo sapiens

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Pro Pro Asp Phe Asp Pro Pro Trp Val Glu Lys Ala Asn Arg Thr Arg 50 55 60

Gly Val Gly Lys Glu Ala Ser Lys Ala Leu Gly Ala Lys Gly Ser Cys 65 70 75 80

Glu Thr Thr Pro Ser Arg Gly Ser Thr Pro Thr Leu Thr Pro Arg Lys
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Lys Asn Lys Tyr Arg Pro Ile Ser His Thr Pro Ser Tyr Cys Asp Glu 100 105 110

Ser	Leu	Phe	Gly	Ser	Arg	Ser	Glu	Gly	Ala	Ser	Phe	Gly	Ala	Pro	Arg
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Pro Pro Thr Pro Arg Gly Ser His Ser Pro Arg Pro Arg Glu Ala Pro 145 150 155 160

Leu Arg Ala Ile His Pro Ala Gly Pro Ser Lys Thr Glu Pro Gly Pro 165 170 175

Ala Ala Asp Ser Gln Lys Leu Ser Met Gly Gly Leu His Ser Ser Arg 180 185 190

Pro Leu Lys Arg Gly Leu Ser His Ser Leu Thr His Leu Asn Val Pro 195 200 205

Ser Thr Gly His Pro Ala Thr Ser Ala Pro His Thr Asn Gly Pro Gln 210 215 220

Asp Leu Arg Pro Ser Thr Ser Gly Val Thr Phe Arg Ser Pro Leu Val 225 230 235 240

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WO 01/07607

11/27

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Pro	Asn	Asn		Leu	Ile	Asn	Asn			Ala	Gly	Ser	61n 45		Ala	
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_	_		_	_	aaa		-									419
Leu	65	Leu	Leu	Ala	Lys	70	ulu	Leu	1mr	Pro	75		116	rne	1 y I	
	00					70					10					
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Asn	Asn	Arg	Val	His	Ser	Asn	Phe	Lys	Ala	Gly	Leu	Phe	Ile	Asp	Lys	
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Gl	y Val	l Lys	s Thi	Thr 100		ser	· Ser	· Ala	105		Pro	Arg	Glu	Tyr 110		
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			Arg	gtt Val												611
		Asp		gga Gly			_						_			659
		_	_	gca Ala												707
	_			agt Ser 180												755
				gag Glu												803
				act Thr		Gly		_	_	_		_				851
				ttc Phe	Pro		_			Gln			_			899
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							gaa Glu									1619
							acg Thr								aaa Lys 495	1667
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											cgt Arg					1955
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											ttc Phe					2051
					Leu.					Arg	att Ile 635					2099
											ttg Leu					2147
_											gta Val		_		_	2195
gcc	aaa	cca	gct	cat	ctt	tat	gac	aaa	ggg	agt	acc	gta	ttt	ttg	gga	2243

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gac gaa tat ggt tgt ccc aga gcc acc act gtc cgc aga aga gac ctg 2387 Asp Glu Tyr Gly Cys Pro Arg Ala Thr Thr Val Arg Arg Arg Asp Leu 720 725 730 735

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PCT/JP00/04895

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Ile Trp Tyr Leu Phe His Lys Glu Pro Thr Gly Glu Ser Ser Gly Leu 50 55 60

Gln Leu Leu Ala Lys Pro Glu Leu Thr Pro Leu Gly Ile Phe Tyr Asn 65 70 75 80

Asn Arg Val His Ser Asn Phe Lys Ala Gly Leu Phe Ile Asp Lys Gly 85 90 95

Val Lys Thr Thr Asn Ser Ser Ala Ala Asp Pro Arg Glu Tyr Leu Cys 100 105 110

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- Asn Arg Thr Phe Pro Ile Arg Gly Phe Gln Ile Tyr Asp Gly Pro Ile

- His Leu Thr Arg Ser Thr Phe Lys Lys Tyr Val Pro Thr Pro Asp Arg
- Tyr Ser Ser Ala Ile Gly Phe Leu Met Lys Asn Ser Trp Gln Ile Thr
- Pro Arg Asn Asn Ile Ser Leu Val Lys Phe Gly Pro His Val Ser Leu
- Asn Val Phe Phe Gly Lys Pro Gly Pro Trp Phe Glu Asp Cys Glu Met

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Ile	Lys	Gln	Leu 660	Asn	Ile	Ser	His	Leu 665	Leu	Val	Pro	Leu	Gly 670	Leu	Ala
Lys	Pro	Ala 675	His	Leu	Tyr	Asp	Lys 680	Gly	Ser	Thr	Val	Phe 685	Leu	Gly	Phe

Ser Gly Asn Phe Lys Pro Ser Trp Thr Lys Leu Phe Thr Ser Pro Ala 690 695 700

Gly Gln Gly Leu Gly Val Leu Glu Gln Phe Ile Pro Leu Gln Leu Asp 705 710 715 720

Glu Tyr Gly Cys Pro Arg Ala Thr Thr Val Arg Arg Arg Asp Leu Glu
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Leu Leu Lys Gln Ala Ser Lys Ala His
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Met Pro Leu Val Gly

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				tgg Trp						451
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_				aac Asn 60						547
				gca Ala						595
			Arg	gtc Val		Ile			Gly	643
				cct Pro						691
				tac Tyr						739
				ctg Leu						787

	135					140					145					
		_												ggc Gly		835
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-														aca Thr		979
Glu			cac His					ctc Leu	tgad	cace	cc g	gece	ectte	ca		1026
agcg	acca	ica o	tcca	ıccat	c to	agac	agca	ı gca	ccto	ectc	ttct	tagca	agc (	cagto	ectect	1086
ccat	cctg	gg. e	tcgc	tggg	c.ct	gctt	gtgt	, cct	ccag	ccc	agco	cac	ccg, į	ggcct	tatgga	1146
gccc	tgcc	ca c	agco	cctg	g to	atct	gata	tca	gago	ctg	cgto	gagg	gaa į	gatga	agccag	1206
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25/27

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Arg Ile Gln Glu Met Ala Ser Met Gly Ile Gly Asn Gln Pro Phe Met 50 55 60

Asp Val Lys Pro Arg Asp Arg Thr Pro Asp Cys Ala Val Ile Ser Asp 65 70 75. 80

Gly Ala Pro Lys Cys Ala Val Met Ser Asp Arg Val Pro Asp Ser Ile 85 90 95

Val Lys Gly Thr Gly Thr Val Ala Arg Ser Arg Pro His Ser Pro Cys 100 105 110

Arg Gly His Trp Ala Cys His Gln Gly His Gly Tyr Gly Gly Ile Gly
115 120 125

Pro Thr Leu Thr Arg Pro Gln Ser Ala Pro Gly Leu Ser Ser Arg Thr

130 135 140

Arg Val Arg Trp Pro Arg Pro Arg Pro His Ser Ser Cys Arg Gly His 145 150 155 160

Trp Ala Ser Gly Arg His Gly Gly Leu Asp Gly His Asp Cys Ser Gly 165 170 175

Lys Ala Trp Ser Ala Phe Gln Thr Ala Leu Ile Pro Phe Pro Asn Leu 180 185 190

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